

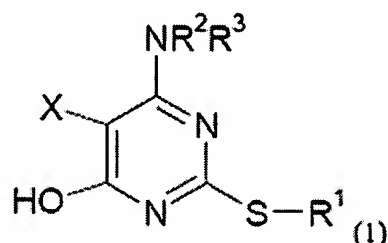
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DT01 Rec'd PCT/PTC 23 FEB 2005

Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

Listing of Claims:

1. (Currently amended) A compound of formula (1), a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof:



wherein R^1 is a group selected from C_{3-7} carbocyclyl, C_{1-8} alkyl, C_{2-6} alkenyl and C_{2-6} alkynyl; wherein the group is optionally substituted by 1, 2 or 3 substituents independently selected from fluoro, nitrile, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, $-\text{NR}^8\text{COR}^9$, $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{NR}^8\text{SO}_2\text{R}^9$, phenyl or heteroaryl; wherein phenyl and heteroaryl are optionally substituted by 1, 2 or 3 substituents independently selected from halo, cyano, nitro, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, $-\text{NR}^8\text{COR}^9$, $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{NR}^8\text{SO}_2\text{R}^9$, C_{1-6} alkyl and trifluoromethyl;

wherein R^2 is C_{3-7} carbocyclyl, optionally substituted by 1, 2 or 3 substituents independently selected from:

- (a) fluoro, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, $-\text{NR}^8\text{COR}^9$, $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{NR}^8\text{SO}_2\text{R}^9$;
- (b) a 3-8 membered ring optionally containing 1, 2 or 3 atoms selected from O, S, $-\text{NR}^8$ and whereby the ring is optionally substituted by C_{1-3} alkyl or fluoro; or

(c) phenyl or heteroaryl, each of which is optionally substituted by 1, 2 or 3 substituents independently selected from halo, cyano, nitro, $-OR^4$, $-NR^5R^6$, $-CONR^5R^6$, $-NR^8COR^9$, $-SO_2NR^5R^6$, $-NR^8SO_2R^9$, C_{1-6} alkyl and trifluoromethyl;

or R^2 is a group selected from C_{1-6} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl wherein the group is substituted by 1, 2 or 3 substituents independently selected from hydroxy, amino, C_{1-6} alkoxy, C_{1-6} alkylamino, di(C_{1-6} alkyl)amino, N -(C_{1-6} alkyl)- N -(phenyl)amino, N - C_{1-6} alkylcarbamoyl, N,N -(C_{1-6} alkyl)₂carbamoyl, N -(C_{1-6} alkyl)- N -(phenyl)carbamoyl, carboxy, phenoxycarbonyl, $-NR^8COR^9$, $-SO_2R^{10}$, $-SO_2NR^5R^6$ and $-NR^8SO_2R^9$;

wherein R^3 is hydrogen or R^2 ;

R^4 is hydrogen or a group selected from C_{1-6} alkyl and phenyl, wherein the group is optionally substituted by 1 or 2 substituents independently selected from halo, phenyl, $-OR^{11}$ and $-NR^{12}R^{13}$;

R^5 and R^6 are independently hydrogen or a group selected from C_{1-6} alkyl and phenyl wherein the group is optionally substituted by 1, 2 or 3 substituents independently selected from halo, phenyl, $-OR^{14}$, $-NR^{15}R^{16}$, $-CONR^{15}R^{16}$, $-NR^{15}COR^{16}$, $-SONR^{15}R^{16}$ and $NR^{15}SO_2R^{16}$

or

R^5 and R^6 together with the nitrogen atom to which they are attached form a 4- to 7-membered saturated heterocyclic ring system optionally containing a further heteroatom selected from oxygen and nitrogen atoms, which ring system may be optionally substituted by 1, 2 or 3 substituents independently selected from phenyl, $-OR^{14}$, $-COOR^{14}$, $-NR^{15}R^{16}$, $-CONR^{15}R^{16}$, $-NR^{15}COR^{16}$, $-SONR^{15}R^{16}$, $NR^{15}SO_2R^{16}$ or C_{1-6} alkyl₂ (optionally substituted by 1 or 2 substituents independently selected from halo, $-NR^{15}R^{16}$ and $-OR^{17}$ groups);

R^{10} is hydrogen or a group selected from C_{1-6} alkyl or phenyl, wherein the group is optionally substituted by 1, 2 or 3 substituents independently selected from halo, phenyl, $-OR^{17}$ and $-NR^{15}R^{16}$;

and

each of R^7 , R^8 , R^9 , R^{11} , R^{12} , R^{13} , R^{14} , R^{15} , R^{16} , R^{17} is independently hydrogen, C_{1-6} alkyl or phenyl;

X is hydrogen, halo, cyano, nitro, hydroxy, C_{1-6} alkoxy₂ (optionally substituted by 1 or 2 substituents selected from halo, $-OR^{11}$ and $-NR^{12}R^{13}$), $-NR^5R^6$, $-COOR^7$, $-CONR^5R^6$, $-NR^8COR^9$, thio, thiocyano, thio C_{1-6} alkyl₂ (optionally substituted by 1 or 2 substituents selected from halo,

$-\text{OR}^{17}$, $-\text{COOR}^7$, $-\text{NR}^{15}\text{R}^{16}$, $-\text{CONR}^5\text{R}^6$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{NR}^8\text{SO}_2\text{R}^{10}$ or a group selected from C_{3-7} carbocyclyl, C_{1-8} alkyl, C_{2-6} alkenyl or C_{2-6} alkynyl, wherein the group is optionally substituted by 1, 2 or 3 substituents independently selected from halo, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, $-\text{NR}^8\text{COR}^9$, $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$ and $-\text{NR}^8\text{SO}_2\text{R}^9$; or a -phenyl, -heteroaryl, -thiophenyl, -thioheteroaryl, aminoheteroaryl, and thio C_{1-6} alkylheteroaryl group, all of which may be optionally substituted by 1, 2 or 3 substituents independently selected from halo, cyano, nitro, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, $-\text{NR}^8\text{COR}^9$, $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{NR}^8\text{SO}_2\text{R}^9$, $\text{C}_1\text{-C}_6$ alkyl, phenyl, heteroaryl or trifluoromethyl groups;

2. (Currently amended) A compound according to claim 1 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof, wherein R^1 is C_{1-8} alkyl optionally substituted by 1, 2 or 3 substituents independently selected from phenyl or heteroaryl, wherein phenyl and heteroaryl are optionally substituted by 1, 2 or 3 substituents independently selected from halo, cyano, $-\text{OR}^4$, $-\text{SR}^{10}$, C_{1-6} alkyl and trifluoromethyl.

3. (Currently amended) A compound according to claim 1 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof, wherein R^2 is C_{1-8} alkyl substituted by 1, 2 or 3 substituents independently selected from hydroxy, amino, C_{1-6} alkoxy, C_{1-6} alkylamino, di(C_{1-6} alkyl)amino, *N*-(C_{1-6} alkyl)-*N*-(phenyl)amino, *N*-(C_{1-6} alkyl)carbamoyl, *N,N*-di(C_{1-6} alkyl)carbamoyl, *N*-(C_{1-6} alkyl)-*N*-(phenyl)carbamoyl, carboxy, phenoxycarbonyl, $-\text{NR}^8\text{COR}^9$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$ and $-\text{NR}^8\text{SO}_2\text{R}^9$; and wherein R^3 is hydrogen;

4. (Currently amended) A compound according to claim 1 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof, wherein R^4 , R^5 , R^6 , R^8 , R^9 and R^{10} are independently hydrogen, C_{1-6} alkyl or phenyl.

5. (Currently amended) A compound according to claim 1 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof, wherein X is hydrogen, halo, cyano, nitro,

hydroxy, thio, thiocyno, $-\text{CONR}^5\text{R}^6$, thio $\text{C}_{1-6}\text{alkyl}_2$ (optionally substituted by 1 or 2 substituents selected from halo, $-\text{OR}^{17}$, $-\text{NR}^{15}\text{R}^{16}$, $-\text{CONR}^5\text{R}^6$), $-\text{NR}^8\text{SO}_2\text{R}^{10}$, $\text{C}_{1-8}\text{alkyl}_2$ (optionally substituted by 1, 2 or 3 substituents independently selected from halo, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, $-\text{NR}^8\text{COR}^9$, $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$ and $-\text{NR}^8\text{SO}_2\text{R}^9$), heteroaryl, thioheteroaryl or thio $\text{C}_{1-6}\text{alkylheteroaryl}$ all of which may be optionally substituted by 1, 2 or 3 substituents independently selected from halo, cyano, nitro, $-\text{OR}^4$, $-\text{NR}^5\text{R}^6$, $-\text{CONR}^5\text{R}^6$, $-\text{COOR}^7$, NR^8COR^9 , $-\text{SR}^{10}$, $-\text{SO}_2\text{R}^{10}$, $-\text{SO}_2\text{NR}^5\text{R}^6$, $-\text{NR}^8\text{SO}_2\text{R}^9$, $\text{C}_{1-6}\text{alkyl}$ or trifluoromethyl.

6. (Currently amended) A compound according to claim 2 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof; wherein R^1 is benzyl optionally substituted by 1 or 2 substituents independently selected from fluoro, chloro, bromo, methoxy, methyl and trifluoromethyl.

7. (Currently amended) A compound according to claim 3 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof; wherein R^2 is $\text{C}_{1-4}\text{alkyl}$, substituted by 1, 2 or 3 substituents independently selected from hydroxy, amino, $\text{C}_{1-6}\text{alkoxy}$, $\text{C}_{1-6}\text{alkylamino}$, and $\text{di}(\text{C}_{1-6}\text{alkyl})\text{amino}$; and R^3 is hydrogen.

8. (Currently amended) A compound according to claim 4 or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof; wherein X is hydrogen, fluoro, chloro, bromo, thiocyno, $-\text{NR}^8\text{SO}_2\text{R}^9$ (where R^8 is hydrogen and R^9 is methyl), $-\text{thioimidazolyl}$, $-\text{thiotriazolyl}$, $-\text{CONH}_2$, $-\text{CONMe}_2$ or cyano.

9. (Original) A compound selected from the group consisting of:

2-(Benzylthio)-6- $\{[(1R)\text{-}2\text{-hydroxy-}1\text{-methylethyl}]\text{amino}\}$ -4-pyrimidinol,

2-(Benzylthio)-5-chloro-6- $\{[(1R)\text{-}2\text{-hydroxy-}1\text{-methylethyl}]\text{amino}\}$ -4-pyrimidinol,

2-[(3-Chlorobenzyl)thio]-6- $\{[(1R)\text{-}2\text{-hydroxy-}1\text{-methylethyl}]\text{amino}\}$ -4-pyrimidinol,

5-Chloro-2-[(3-chlorobenzyl)thio]-6- $\{[(1R)\text{-}2\text{-hydroxy-}1\text{-methylethyl}]\text{amino}\}$ -4-pyrimidinol,

2-[(3-Chlorobenzyl)thio]-4-hydroxy-6-[(1R)-2-hydroxy-1-methylethyl]amino}-5-pyrimidinyl
thiocyanate,
N-(2-[(3-Chlorobenzyl)thio]-4-hydroxy-6-[(1R)-2-hydroxy-1-methylethyl]amino)-5-
pyrimidinyl)methanesulfonamide,
2-[(3-Chlorobenzyl)thio]-5-fluoro-6-[(1R)-2-hydroxy-1-methylethyl]amino}-4-pyrimidinol, 2-
[(2,3-difluorobenzyl)thio]-4-hydroxy-6-[(1S)-2-hydroxy-1-methylethyl]amino}pyrimidine-5-
carbonitrile,
5-Chloro-2-[(2,3-difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-4-
pyrimidinol,
2-[(2,3-Difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-iodo-4-
pyrimidinol,
2-[(2,3-Difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-nitro-4-
pyrimidinol,
2-[(3-Chlorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-(1,3,4-
thiadiazol-2-ylthio)-4-pyrimidinol,
2-[(2,3-Difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-(1*H*-
imidazol-2-ylthio)-4-pyrimidinol,
2-[(2,3-Difluorophenyl)methyl]thio]-5-[2-(dimethylamino)ethyl]thio]-6-[(1R)-2-hydroxy-1-
methylethyl]amino]-4-pyrimidinol,
1-[2-[(2,3-Difluorophenyl)methyl]thio]-4-hydroxy-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-
pyrimidinyl)- 4(1*H*)-pyridinethione,
2-[(2,3-Difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-(4-
pyridinylthio)- 4-pyrimidinol,
2-[(2,3-Difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-(1*H*-1,2,4-
triazol-3-ylthio)- 4-pyrimidinol,
2-[(2,3-Difluorophenyl)methyl]thio]-6-[(1R)-2-hydroxy-1-methylethyl]amino]-5-[(4-methyl-
4*H*-1,2,4-triazol-3-yl)thio]- 4-pyrimidinol,
5-[(5-Amino-4*H*-1,2,4-triazol-3-yl)thio]-2-[(2,3-difluorophenyl)methyl]thio]-6-[(1R)-2-

hydroxy-1-methylethyl]amino]- 4-pyrimidinol,
2-[[[(2,3-Difluorophenyl)methyl]thio]-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-[[5-(4-pyridinyl)-1,3,4-oxadiazol-2-yl]thio]- 4-pyrimidinol,
Ethyl[[2-[[[(2,3-difluorophenyl)methyl]thio]-4-hydroxy-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-pyrimidinyl]thio]- AcOH,
2-[[2-[[[(2,3-Difluorophenyl)methyl]thio]-4-hydroxy-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-pyrimidinyl]thio]-*N*-methyl- acetamide,
2-[[2-[[[(2,3-Difluorophenyl)methyl]thio]-4-hydroxy-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-pyrimidinyl]thio]-*N*-[2-(dimethylamino)ethyl]- acetamide,
1-[[[2-[[[(2,3-Difluorophenyl)methyl]thio]-4-hydroxy-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-pyrimidinyl]thio]acetyl]-piperazine,
2-[[[(2,3-Difluorophenyl)methyl]thio]-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-[(4-methyl-2-oxazolyl)thio]- 4-pyrimidinol,
2-[[[(2,3-Difluorophenyl)methyl]thio]-6-[[[(1*R*)-2-hydroxy-1-methylethyl]amino]-5-[(1,2,4-oxadiazol-3-ylmethyl)thio]- 4-pyrimidinol,
2-[(2,3-difluorobenzyl)thio]-4- {[[(1*R*)-1,2-dihydroxyethyl]amino}-6-hydroxypyrimidine-5-carboxamide,
2-[(2,3-difluorobenzyl)thio]-6- {[[(1*R*)-2-hydroxy-1-methylethyl]amino}-5-(5-methyl-1,2,4-oxadiazol-3-yl)pyrimidin-4-ol,
2-[(2,3-difluorobenzyl)thio]-6- {[[(1*R*)-2-hydroxy-1-methylethyl]amino}-5-(1,3-oxazol-5-yl)pyrimidin-4-ol,
2-[(2,3-difluorobenzyl)thio]-4- {[[(1*R*)-1,2-dihydroxyethyl]amino}-6-hydroxy-*N,N*-dimethylpyrimidine-5-carboxamide,
2-[(2,3-difluorobenzyl)thio]-5-fluoro-6- {[[(1*R*)-2-hydroxy-1-methylethyl]amino}-pyrimidin-4-ol,
2-[(3,4-difluorobenzyl)thio]-5-fluoro-6- {[[(1*R*)-2-hydroxy-1-methylethyl]amino}-pyrimidin-4-ol,
2-[(3-fluorobenzyl)thio]-5-fluoro-6- {[[(1*R*)-2-hydroxy-1-methylethyl]amino}pyrimidin-4-ol,
or
2-[(4-fluorobenzyl)thio]-5-fluoro-6- {[[(1*R*)-2-hydroxy-1-methylethyl]amino}pyrimidin-4-ol and

a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof.

10. (Cancelled)

11. (Currently amended) A method of treating, compound, pharmaceutically acceptable salt, solvate, or in vivo hydrolysable ester thereof according to any one of claims 1 to 9 for use as a medicament for the treatment of asthma, allergic rhinitis, COPD, inflammatory bowel disease, osteoarthritis, osteoporosis, rheumatoid arthritis, or psoriasis comprising administering a compound of claim 1, or a pharmaceutically acceptable salt, solvate or in vivo hydrolysable ester thereof.

12. (Currently amended) A method of treating cancer, comprising administering a compound of claim 1, or a pharmaceutically acceptable salt, solvate or in vivo hydrolysable ester thereof. ; ~~pharmaceutically acceptable salt, solvate, or in vivo hydrolysable ester thereof according to any one of claims 1 to 9 for use as a medicament for the treatment of cancer.~~

13. (Currently amended) A method of treating COPD comprising administering a compound of claim 1, or a pharmaceutically acceptable salt, solvate or in vivo hydrolysable ester thereof.; ~~pharmaceutically acceptable salt, solvate, or in vivo hydrolysable ester thereof according to any one of claims 1 to 9 for use as a medicament for the treatment of COPD.~~

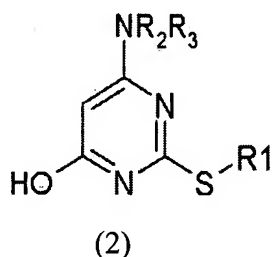
14. (Currently amended) A method of treating a disease or condition The use of a compound, pharmaceutically acceptable salt, solvate, or in vivo hydrolysable ester thereof according to any one of claims 1 to 9 in the manufacture of a medicament for the treatment of human diseases or conditions in which modulation of chemokine receptor activity is beneficial, comprising administering a compound of claim 1, or a pharmaceutically acceptable salt, solvate or in vivo hydrolysable ester thereof.

15-16. (Cancelled)

17. (Currently amended) A pharmaceutical composition comprising a compound, pharmaceutically acceptable salt, solvate, or in vivo hydrolysable ester thereof according to ~~any one of claims 1 to 9~~ claim 1; and a pharmaceutically-acceptable diluent or carrier.

18. (Currently amended) A process for the preparation of a compound of formula (1) as defined above which comprises

(a) treating a compound of formula (2):

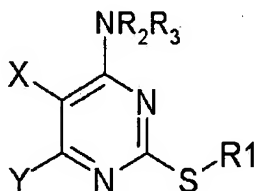


wherein R¹, R² and R³ are as defined in claim 1, formula (1), with suitable electrophiles.

and optionally thereafter (i), (ii), (iii), (iv) or (v) in any order:

- i) removing any protecting groups;
- ii) converting the compound of claim 1, formula (1) into a further compound of claim 1, formula (1),
- iii) forming a salt;
- (iv) forming a prodrug
- (v) forming an in vivo hydrolysable ester; or

(b) , where X is 1,3-oxazol-5-yl by treating a compound of formula (4):



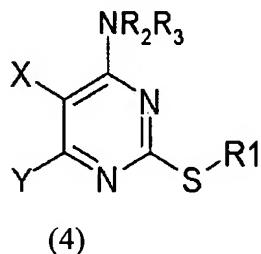
(4)

wherein R^1 , R^2 and R^3 are as defined in claim 1, formula (1), X is $-CHO$ and Y is protected hydroxy by treatment with *p*-toluenesulfonylmethyl isocyanide and potassium hydroxide in refluxing methanol.

and optionally thereafter (i), (ii), (iii), (iv) or (v) in any order:

- i) removing any protecting groups;
- ii) converting the compound of claim 1, formula (1) into a further compound of claim 1, formula (1),
- iii) forming a salt;
- iv) forming a prodrug,
- v) forming an *in vivo* hydrolysable ester; or

(c) where X is CN by treating a compound of formula (4):



wherein R^1 , R^2 and R^3 are as defined in claim 1, formula (1), X is CN and Y is halogen by treatment

with potassium *tert*-butoxide in refluxing aqueous toluene.

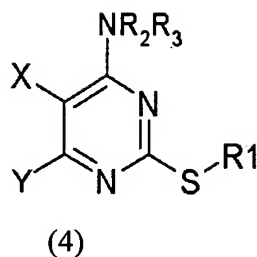
and optionally thereafter (i), (ii), (iii), (iv) or (v) in any order:

- i) removing any protecting groups,
- ii) converting the compound of claim 1, formula (1) into a further compound of claim 1, formula (1),
- iii) forming a salt;

- (iv) forming a prodrug,
- v) forming an *in vivo* hydrolysable ester; or

(d) where X is $-\text{CONR}^5\text{R}^6$ by;

e) treating a compound of formula (4):



wherein R^1 , R^2 and R^3 are as defined in claim 1, formula (1), X is $-\text{CONR}^5\text{R}^6$ and Y is halogen with a suitable base.

and optionally thereafter (i), (ii), (iii), (iv) or (v) in any order:

- i) removing any protecting groups;
- ii) converting the compound of claim 1, formula (1) into a further compound of claim 1, formula (1),
- iii) forming a salt;
- (iv) forming a prodrug,
- v) forming an *in vivo* hydrolysable ester.

19. (Currently amended) A ~~combination therapy~~ method of combination therapy which comprises administering a compound of claim 1, formula (1) or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester thereof, or a pharmaceutical composition or formulation comprising a compound of claim 1, formula (1), concurrently or sequentially with other therapy and/or another pharmaceutical agent.

20. (Currently amended) ~~A combination therapy as claimed in~~ The method of claim 19,
comprising treating for the treatment of asthma, allergic rhinitis, COPD, inflammatory bowel
disease, irritable bowel syndrome, osteoarthritis, osteoporosis, rheumatoid arthritis, or psoriasis.

21. (Currently amended) ~~A combination therapy as claimed in~~ The method of claim 19,
comprising treating for the treatment of cancer.

22. (Currently amended) A pharmaceutical composition which comprises a compound of
claim 1, formula (1) or a pharmaceutically acceptable salt, solvate or *in vivo* hydrolysable ester
thereof, in conjunction with another pharmaceutical agent.

23. (Currently amended) A method of treating pharmaceutical composition as claimed in
~~claim 22 for the treatment of~~ asthma, allergic rhinitis, COPD, inflammatory bowel disease,
irritable bowel syndrome, osteoarthritis, osteoporosis, rheumatoid arthritis, or psoriasis,
comprising administering the pharmaceutical composition of claim 22.

24. (Currently amended) A method of treating pharmaceutical composition as claimed in
~~claim 22 for the treatment of~~ cancer, comprising administering the pharmaceutical composition
of claim 22.